

### AMENDMENTS TO THE CLAIMS

Claim 1 (currently amended): A method for estimating a virtual patient's fasting plasma glucose (FPG) level, comprising:

- determining the virtual patient's basal hepatic production ( $FPG_0$ );
- determining the virtual patient's insulin level ( $I$ );
- calculating the virtual patient's FPG at time  $t$  by solving an equation

$$FPG(t) = FPG_0 / (I * E), \text{ wherein } E \text{ is a value representing efficiency of insulin use;}$$

and

~~saving~~ outputting at least one value ~~based on~~ for the virtual patient's FPG at time  $t$  ~~in a~~  
~~computer-readable medium~~ to a user.

Claim 2 (previously presented): The method of claim 1, wherein  $E$  is scaled such that  $E = 1$  in the absence of diabetes and  $0 \leq E < 1$  in the presence of diabetes.

Claim 3 (currently amended): The method of claim 1, wherein for type 2 diabetes, an equation representing  $E$  is:

$$E(DF_2) = \left( a + b / \left( 1 + (DF_2 / c)^d \right) \right)^{\frac{1}{2}}, \text{ wherein the parameters } a, b, c, \text{ and } d \text{ are set to reflect fit}$$

data for a population that is represented by the virtual patient, and  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.

Claim 4 (previously presented): The method of claim 3, wherein

$$DF_2(t) = \left( 1 - \exp \left( -a * IGT(\xi_3) / \left( 1 + \exp \left( -\frac{(t-b)}{c} \right) \right) \right) \right) * RBMI(BMI) / \xi_2, \text{ wherein } IGT \text{ is an}$$

impaired glucose tolerance value,  $RBMI$  is a relative risk associated with the virtual patient's body mass index ( $BMI$ ), and  $\xi_1$  and  $\xi_3$  are random values selected from distributions for randomizing the virtual patient within the population.

Claim 5 (original): The method of claim 4, wherein the  $RBMI$  is represented by:

$$RBMI(BMI) = a + b / \left( 1 + e^{-(BMI-c)/d} \right).$$

Claim 6 (original): The method of claim 4, wherein  $IGT$  is represented by:

$$IGT(\xi_3) = 2(1 - \xi_3)$$

wherein  $\xi_3$  is a random value designed to cause the occurrence of diabetes in virtual patients to have the same types of interpersonal variations that occur in real people.

Claim 7 (previously presented): The method of claim 1, wherein said determining said virtual patient's basal hepatic production in type 2 diabetes includes solving an equation  $FPG_0(t) = G(t) * H(DF_2(t))$ , wherein  $G(t)$  represent a basal production in people who do not have diabetes,  $H$  represents a degree of insulin resistance in a person with diabetes, and  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.

Claim 8 (currently amended): The method of claim 7, wherein

$H(DF_2(t)) = 1 / \left( \text{MAX} \left[ E^2(DF_2(t+a)), b \right] \right)$ , and the parameters  $a$  and  $b$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claim 9 (currently amended): The method of claim 7, wherein

$G(t) = (a + bt^{1.5} - c * t^3 + \Delta_g) / (d - e \exp(-DF_2(t)\xi_2))$ , wherein  $\Delta_g$  represents a variance of basal hepatic production across individuals, the parameters  $a$ ,  $b$ ,  $c$ ,  $d$ , and  $e$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient, and  $\xi_2$  is a random value selected from a distribution for randomizing the virtual patient within the population.

Claim 10 (currently amended): The method of claim 1, wherein

$I(DF_1, DF_2) = H(DF_2) * E(DF_2) / (1 + \exp((DF_1 - a)/b))$ , and wherein  $DF_1$  is a type 1 diabetes feature that represents an incidence of type 1 diabetes for the virtual patient,  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient,  $H$  represents a degree of insulin resistance in a person with diabetes, and the parameters  $a$  and  $b$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claims 11-30 (cancelled).

Claim 31 (currently amended): An apparatus for estimating a virtual patient's fasting plasma glucose (FPG) level, the apparatus comprising:

means for determining the virtual patient's basal hepatic production ( $FPG_0$ );

means for determining the virtual patient's insulin level ( $I$ );

means for calculating the virtual patient's FPG at time  $t$  by solving an equation

$$FPG(t) = FPG_0 / (I * E), \text{ wherein } E \text{ is a value representing efficiency of insulin use;}$$

and

means for ~~saving~~ outputting at least one value ~~based on~~ for the virtual patient's FPG at time  $t$  to a user.

Claim 32 (previously presented): The apparatus of claim 31, wherein  $E$  is scaled such that  $E = 1$  in the absence of diabetes and  $0 \leq E < 1$  in the presence of diabetes.

Claim 33 (currently amended): The apparatus of claim 31, wherein for type 2 diabetes, an equation representing  $E$  is:

$$E(DF_2) = \left( a + b / \left( 1 + (DF_2 / c)^d \right) \right)^{\frac{1}{2}}, \text{ wherein the parameters } a, b, c, \text{ and } d \text{ are set to } \del{reflect} \u{fit}$$

data for a population that is represented by the virtual patient, and  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.

Claim 34 (previously presented): The apparatus of claim 33,

$$\text{wherein } DF_2(t) = \left( 1 - \exp \left( -a * IGT(\xi_3) / \left( 1 + \exp \left( -\frac{(t-b)}{c} \right) \right) \right) \right) * RBMI(BMI) / \xi_2,$$

wherein  $IGT$  is an impaired glucose tolerance value,  $RBMI$  is a relative risk associated with the virtual patient's body mass index (BMI), and  $\xi_1$  and  $\xi_3$  are random values selected from distributions for randomizing the virtual patient within the population.

Claim 35 (previously presented): The apparatus of claim 33, wherein the *RBMI* is represented by:

$$RBMI(BMI) = a + b / (1 + e^{-(BMI-c)/d}).$$

Claim 36 (previously presented): The apparatus of claim 34, wherein *IGT* is represented by:

$$IGT(\xi_3) = 2(1 - \xi_3),$$

wherein  $\xi_3$  is a random value designed to cause the occurrence of diabetes in virtual patients to have the same types of interpersonal variations that occur in real people.

Claim 37 (previously presented): The apparatus of claim 31, wherein said means for determining said virtual patient's basal hepatic production in type 2 diabetes includes means for solving an equation  $FPG_0(t) = G(t) * H(DF_2(t))$ , wherein  $G(t)$  represent a basal production in people who do not have diabetes,  $H$  represents a degree of insulin resistance in a person with diabetes, and  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.

Claim 38 (currently amended): The apparatus of claim 37, wherein

$H(DF_2(t)) = 1 / (MAX[E^2(DF_2(t+a)), b])$ , and the parameters  $a$  and  $b$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claim 39 (currently amended): The apparatus of claim 37, wherein

$G(t) = (a + bt^{1.5} - c * t^3 + \Delta_g) / (d - e \exp(-DF_2(t)\xi_2))$ , wherein  $\Delta_g$  represents a variance of basal hepatic production across individuals, the parameters  $a$ ,  $b$ ,  $c$ ,  $d$ , and  $e$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient, and  $\xi_2$  is a random value selected from a distribution for randomizing the virtual patient within the population.

Claim 40 (currently amended): The apparatus of claim 31, wherein

$I(DF_1, DF_2) = H(DF_2) * E(DF_2) / (1 + \exp((DF_1 - a)/b))$ , and wherein  $DF_1$  is a type 1 diabetes

feature that represents an incidence of type 1 diabetes for the virtual patient,  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient,  $H$  represents a degree of insulin resistance in a person with diabetes, and the parameters  $a$  and  $b$  are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claims 41-51 (cancelled).

Claim 52 (currently amended): A program storage device readable by a machine, tangibly embodying a program of instructions executable by the machine to perform a method for estimating a virtual patient's fasting plasma glucose (FPG) level, the method comprising:

determining the virtual patient's basal hepatic production ( $FPG_0$ );

determining the virtual patient's insulin level ( $I$ );

calculating the virtual patient's FPG at time  $t$  by solving an equation

$$FPG(t) = FPG_0 / (I * E), \text{ wherein } E \text{ is a value representing efficiency of insulin use;}$$

and

~~saving~~ outputting at least one value ~~based on~~ for the virtual patient's FPG at time  $t$  to a user.

Claims 53-60 (cancelled).

Claim 61 (previously presented): The method of claim 1, wherein the at least one value based on the virtual patient's FPG at time  $t$  is saved in at least one file in a computer storage device.

Claim 62 (previously presented): The method of claim 3, further comprising:

setting values for the parameters  $a$ ,  $b$ ,  $c$ , and  $d$  by fitting the equation representing  $E$  to data for the population according to a least-squares criterion.

Claim 63 (previously presented): The apparatus of claim 31, wherein the at least one value based on the virtual patient's FPG at time  $t$  is saved to a computer-readable medium.

Claim 64 (currently amended): The apparatus of claim ~~34~~ 33, further comprising:

means for setting values for the parameters  $a$ ,  $b$ ,  $c$ , and  $d$  by fitting the equation representing  $E$  to data for the population according to a least-squares criterion.

Claim 65 (previously presented): The program storage device of claim 52, wherein  $E$  is scaled such that  $E = 1$  in the absence of diabetes and  $0 \leq E < 1$  in the presence of diabetes.

Claim 66 (currently amended): The program storage device of claim 52, wherein for type 2 diabetes, an equation representing  $E$  is:

$$E(DF_2) = \left( a + b / \left( 1 + (DF_2 / c)^d \right) \right)^{\frac{1}{2}}, \text{ wherein the parameters } a, b, c, \text{ and } d \text{ are set to reflect fit data for a population that is represented by the virtual patient, and } DF_2 \text{ is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.}$$

Claim 67 (previously presented): The program storage device of claim 66, wherein

$$DF_2(t) = \left( 1 - \exp \left( -a * IGT(\xi_3) / \left( 1 + \exp \left( -\frac{(t-b)}{c} \right) \right) \right) \right) * RBMI(BMI) / \xi_2, \text{ wherein } IGT \text{ is an}$$

impaired glucose tolerance value,  $RBMI$  is a relative risk associated with the virtual patient's body mass index ( $BMI$ ), and  $\xi_1$  and  $\xi_3$  are random values selected from distributions for randomizing the virtual patient within the population.

Claim 68 (previously presented): The program storage device of claim 67, wherein the  $RBMI$  is represented by:

$$RBMI(BMI) = a + b / \left( 1 + e^{-(BMI-c)/d} \right).$$

Claim 69 (previously presented): The program storage device of claim 67, wherein  $IGT$  is represented by:

$$IGT(\xi_3) = 2(1 - \xi_3)$$

wherein  $\xi_3$  is a random value designed to cause the occurrence of diabetes in virtual patients to have the same types of interpersonal variations that occur in real people.

Claim 70 (previously presented): The program storage device of claim 52, wherein said determining said virtual patient's basal hepatic production in type 2 diabetes includes solving an equation  $FPG_0(t) = G(t) * H(DF_2(t))$ , wherein  $G(t)$  represent a basal production in people who do

not have diabetes, H represents a degree of insulin resistance in a person with diabetes, and  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient.

Claim 71 (currently amended): The program storage device of claim 70, ~~The method of claim 7,~~ wherein  $H(DF_2(t)) = 1 / (\text{MAX}[E^2(DF_2(t+a)), b])$ , and the parameters a and b are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claim 72 (currently amended): The program storage device of claim 70, wherein  $G(t) = (a + bt^{1.5} - c * t^3 + \Delta_g) / (d - e \exp(-DF_2(t)\xi_2))$ , wherein  $\Delta_g$  represents a variance of basal hepatic production across individuals, the parameters a, b, c, d, and e are set to ~~reflect~~ fit data for a population that is represented by the virtual patient, and  $\xi_2$  is a random value selected from a distribution for randomizing the virtual patient within the population.

Claim 73 (currently amended): The program storage device of claim 52, wherein  $I(DF_1, DF_2) = H(DF_2) * E(DF_2) / (1 + \exp((DF_1 - a)/b))$ , and wherein  $DF_1$  is a type 1 diabetes feature that represents an incidence of type 1 diabetes for the virtual patient,  $DF_2$  is a type 2 diabetes feature that represents an incidence of type 2 diabetes for the virtual patient, H represents a degree of insulin resistance in a person with diabetes, and the parameters a and b are set to ~~reflect~~ fit data for a population that is represented by the virtual patient.

Claim 74 (previously presented): The program storage device of claim 52, wherein the at least one value based on the virtual patient's FPG at time t is saved to a computer-readable medium.

Claim 75 (currently amended): The program storage device of claim ~~52~~ 66, wherein the method further comprises:

setting values for the parameters a, b, c, and d by fitting the equation representing E to data for the population according to a least-squares criterion.